

<110> Ghayur, Tarig et al.

<130> BBI-149

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<150> 60/181,608

<151> 2000-02-10

<160> 71

<170> PatentIn Ver. 2.1

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Tyr Phe Gly Lys Leu Glu Ser Lys Leu Ser Val Ile Arg Asn Leu Asn
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Asp Gln Val Leu Phe Ile Asp Gln Gly Asn Arg Pro Leu Phe Glu Asp
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Met Thr Asp Ser Asp Cys Arg Asp Asn Ala Pro Arg Thr Ile Phe Ile
35 40 45

Ile Ser Met Tyr Lys Asp Ser Gln Pro Arg Gly Met Ala Val Thr Ile
50 55 60

Ser Val Lys Cys Glu Lys Ile Ser Thr Leu Ser Cys Glu Asn Lys Ile
65 70 75 80

Ile Ser Phe Lys Glu Met Asn Pro Pro Asp Asn Ile Lys Asp Thr Lys
85 90 95

Ser Asp Ile Ile Phe Phe Gln Arg Ser Val Pro Gly His Asp Asn Lys
100 105 110

Met Gln Phe Glu Ser Ser Ser Tyr Glu Gly Tyr Phe Leu Ala Cys Glu
115 120 125

Lys Glu Arg Asp Leu Phe Lys Leu Ile Leu Lys Lys Glu Asp Glu Leu
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Gly Asp Arg Ser Ile Met Phe Thr Val Gln Asn Glu Asp
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Gly Gln Asp Met Glu Gln Gln Val Val Phe Ser Met Ser Phe Val Gln
35 40 45

Gly Glu Glu Ser Asn Asp Lys Ile Pro Val Ala Leu Gly Leu Lys Glu
50 55 60

Lys Asn Leu Tyr Leu Ser Cys Val Leu Lys Asp Asp Lys Pro Thr Leu
65 70 75 80

Gln Leu Glu Ser Val Asp Pro Lys Asn Tyr Pro Lys Lys Lys Met Glu
85 90 95

Lys Arg Phe Val Phe Asn Lys Ile Glu Ile Asn Asn Lys Leu Glu Phe
100 105 110

Glu Ser Ala Gln Phe Pro Asn Trp Tyr Ile Ser Thr Ser Gln Ala Glu
115 120 125

Asn Met Pro Val Phe Leu Gly Gly Thr Lys Gly Gly Gln Asp Ile Thr
130 135 140

Asp Phe Thr Met Gln Phe Val Ser Ser
145 150

T05029"55029"60

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			20					25					30			
Asn	Val	Asn	Leu	Glu	Glu	Lys	Ile	Asp	Val	Val	Pro	Ile	Glu	Pro	His	
		35					40					45				
Ala	Leu	Phe	Leu	Gly	Ile	His	Gly	Gly	Lys	Met	Cys	Leu	Ser	Cys	Val	
	50					55					60					
Lys	Ser	Gly	Asp	Glu	Thr	Arg	Leu	Gln	Leu	Glu	Ala	Val	Asn	Ile	Thr	
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Asp	Leu	Ser	Glu	Asn	Arg	Lys	Gln	Asp	Lys	Arg	Phe	Ala	Phe	Ile	Arg	
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Ser	Asp	Ser	Gly	Pro	Thr	Thr	Ser	Phe	Glu	Ser	Ala	Ala	Cys	Pro	Gly	
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Trp	Phe	Leu	Cys	Thr	Ala	Met	Glu	Ala	Asp	Gln	Pro	Val	Ser	Leu	Thr	
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Lys Ser Trp Tyr Lys Ser Ser Gly Ser Gln Glu His Val Glu Leu Asn
          35             40             45
Pro Arg Ser Ser Ser Arg Ile Ala Leu His Asp Cys Val Leu Glu Phe
          50             55             60
Trp Pro Val Glu Leu Asn Asp Thr Gly Ser Tyr Phe Phe Gln Met Lys
  65             70             75             80
Asn Tyr Thr Gln Lys Trp Lys Leu Asn Val Ile Arg Arg Asn Lys His

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Ile Thr Trp Tyr Lys Asp Asp Ser Lys Thr Pro Val Ser Thr Glu Gln
          35           40           45
Ala Ser Arg Ile His Gln His Lys Glu Lys Leu Trp Phe Val Pro Ala
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Lys Val Glu Asp Ser Gly His Tyr Tyr Cys Val Val Arg Asn Ser Ser

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Gly	Lys	Asn	Asn	Arg	Pro	Ser
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Gly Asn Asp Gln Arg Pro
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Ala Ile Ser Gly Ser Gln Gly Ser Thr Tyr Tyr Ala Asp Ser Val Lys
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Gly

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Ala Ile Ser Gly Ser Gly Gly Ser Thr Trp Tyr Ala Asp Ser Val Lys
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Gly

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Phe Thr Phe Ser Ser Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly

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Lys Gly Leu Glu Trp Val Ser Ala Ile Ser Gly Ser Gly Gly Ser Thr
35 40 45

Tyr Tyr Ala Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn
50 55 60

Ser Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp
65 70 75 80

Thr Ala Val Tyr Tyr Cys Ala Arg Asp Asp Asp Tyr Asp Phe Asp
85 90 95

Tyr Trp Gly Arg Gly Thr Met Val Thr Val Ser Ser
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20 25 30

Ala Val Asn Trp Tyr Gln Gln Leu Pro Gly Thr Ala Pro Lys Leu Leu
35 40 45

Ile Tyr Gly Asn Asp Gln Arg Pro Ser Gly Val Pro Asp Arg Phe Ser
50 55 60

Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Ser Gly Leu Gln
65 70 75 80

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85 90 95

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100 105 110

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20 25 30

Lys Gly Leu Glu Trp Val Ser Ala Ile Ser Gly Ser Gly Gly Ser Thr
35 40 45

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Tyr Tyr Ala Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn
50 55 60

Ser Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp
65 70 75 80

Thr Ala Val Tyr Tyr Cys Ala Arg Asp Asp Asp Asp Tyr Asp Phe Asp
85 90 95

Tyr Trp Gly Arg Gly Thr Met Val Thr Val Ser Ser Gly Gly Gly Gly
100 105 110

Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Ala Gln Ser Val Leu
115 120 125

Thr Gln Pro Pro Ser Ala Ser Gly Thr Pro Gly Gln Arg Val Thr Ile
130 135 140

Ser Cys Ser Gly Ser Ser Ser Asn Ile Gly Ile Asn Ala Val Asn Trp
145 150 155 160

Tyr Gln Gln Leu Pro Gly Thr Ala Pro Lys Leu Leu Ile Tyr Gly Asn
165 170 175

Asp Gln Arg Pro Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Lys Ser
180 185 190

Gly Thr Ser Ala Ser Leu Ala Ile Ser Gly Leu Gln Ser Glu Asp Glu
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Ala Asp Tyr Tyr Cys Ala Ala Trp Asp Asp Ser Leu Ser Gly Pro Val
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Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly
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1 5 10

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Met Tyr Lys Asp Ser Gln Pro Arg Gly Met Ala Val Thr Ile
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Gln Pro Arg Gly Met Ala Val Thr Ile Ser Val Lys Cys Glu
1 5 10

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Ala Val Thr Ile Ser Val Lys Cys Glu Lys Ile Ser Thr Leu
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Asn Lys Met Gln Phe Glu Ser Ser Ser Tyr Glu Gly Tyr Phe
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Glu Gly Tyr Phe Leu Ala Cys Glu Lys Glu Arg Asp Leu Phe
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Ala Cys Glu Lys Glu Arg Asp Leu Phe Lys Leu Ile Leu Lys
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Met Gln Phe Glu Ser Ser Ser Tyr Glu Gly Tyr Phe Leu Ala Cys Glu
115 120 125

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20 25 30

Tyr Ile His Trp Val Arg Gln Ala Pro Gly Gln Gly Phe Glu Trp Ile
 35 40 45
 Gly Arg Leu Asn Pro Thr Thr Gly Asp Ala Asn Phe Ala Glu Lys Phe
 50 55 60
 Gln Gly Arg Val Ala Leu Thr Arg Asp Thr Ser Ile Ser Thr Ala Tyr
 65 70 75 80
 Leu Gln Leu Asp Ser Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Gly Lys Glu Gly Ala Trp Gly Gln Gly Thr Leu Val Thr Val Ser
 100 105 110
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 Thr Val Arg Ile Thr Cys Gln Gly Asp Ser Leu Arg His Phe Tyr Pro
 20 25 30
 aac tgg tac cag cag aag cca gga cag gcc cct gta ctt gtc atc tat 144
 Asn Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro Val Leu Val Ile Tyr
 35 40 45
 ggt aaa aac aat cgg ccc tca ggg atc cca gac cga ttc tct ggc tcc 192
 Gly Lys Asn Asn Arg Pro Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
 50 55 60
 ggc tca gga aac aca ggt tcc ttg acc atc act ggg gcc cag gcg gaa 240
 Gly Ser Gly Asn Thr Gly Ser Leu Thr Ile Thr Gly Ala Gln Ala Glu
 65 70 75 80
 gat gag gct gac tat tac tgt ggc tcc cgg gac agc agt ggt atc cat 288
 Asp Glu Ala Asp Tyr Tyr Cys Gly Ser Arg Asp Ser Ser Gly Ile His
 85 90 95
 gtg gta ttc ggc gga ggg acc aag gtc acc gtc cta ggt 327
 Val Val Phe Gly Gly Gly Thr Lys Val Thr Val Leu Gly
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tcc	ctg	aga	ctc	tcc	tgt	gca	gcc	tct	gga	ttc	acc	ttt	agc	agc	tat	96
Ser	Leu	Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe	Thr	Phe	Ser	Ser	Tyr	
			20					25					30			
gcc	atg	agc	tgg	gtc	cgc	cag	gct	cca	ggg	aag	ggg	ctg	gag	tgg	gtc	144
Ala	Met	Ser	Trp	Val	Arg	Gln	Ala	Pro	Gly	Lys	Gly	Leu	Glu	Trp	Val	
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tca	gct	att	agt	ggc	agt	ggc	ggc	agc	aca	tac	tac	gca	gac	tcc	gtg	192
Ser	Ala	Ile	Ser	Gly	Ser	Gly	Gly	Ser	Thr	Tyr	Tyr	Ala	Asp	Ser	Val	
	50					55					60					
aag	ggc	cgg	ttc	acc	atc	tcc	aga	gac	aat	tcc	aag	aac	acg	ctg	tat	240
Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ser	Lys	Asn	Thr	Leu	Tyr	
65					70					75					80	
ctg	caa	atg	aac	agc	ctg	aga	gcc	gag	gac	acg	gcc	gtg	tat	tac	tgt	288
Leu	Gln	Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr	Ala	Val	Tyr	Tyr	Cys	
				85					90					95		
gcg	aga	gat	gac	gat	gac	tac	gac	ttt	gac	tac	tgg	ggc	cgg	ggg	aca	336
Ala	Arg	Asp	Asp	Asp	Asp	Tyr	Asp	Phe	Asp	Tyr	Trp	Gly	Arg	Gly	Thr	

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Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val			
35 40 45			
Ser Ala Ile Ser Gly Ser Gly Gly Ser Thr Tyr Tyr Ala Asp Ser Val			
50 55 60			
Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ser Lys Asn Thr Leu Tyr			
65 70 75 80			
Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys			
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1 5 10 15			
agg gtc acc atc tct tgt tct gga agc agc tcc aac atc gga att aat			96
Arg Val Thr Ile Ser Cys Ser Gly Ser Ser Ser Asn Ile Gly Ile Asn			
20 25 30			
gct gta aac tgg tac cag cag ctc cca gga acg gcc ccc aaa ctc ctc			144
Ala Val Asn Trp Tyr Gln Gln Leu Pro Gly Thr Ala Pro Lys Leu Leu			
35 40 45			
atc tat ggt aat gat cag cgg ccc tca ggg gtc cct gac cga ttc tct			192
Ile Tyr Gly Asn Asp Gln Arg Pro Ser Gly Val Pro Asp Arg Phe Ser			

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ggc tcc aag tct ggc acc tca gcc tcc ctg gcc atc agt ggg ctc cag			240
Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Ser Gly Leu Gln			
65	70	75	80
tct gag gat gag gct gat tat aac tgt gca gca tgg gat gac agc ctg			288
Ser Glu Asp Glu Ala Asp Tyr Asn Cys Ala Ala Trp Asp Asp Ser Leu			
	85	90	95
agt ggt ccg gtg ttc ggc gga ggg acc aag ctg acc gtc cta ggt g			334
Ser Gly Pro Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly			
	100	105	110

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			20					25					30		
Ala	Val	Asn	Trp	Tyr	Gln	Gln	Leu	Pro	Gly	Thr	Ala	Pro	Lys	Leu	Leu
		35					40					45			
Ile	Tyr	Gly	Asn	Asp	Gln	Arg	Pro	Ser	Gly	Val	Pro	Asp	Arg	Phe	Ser
	50					55					60				
Gly	Ser	Lys	Ser	Gly	Thr	Ser	Ala	Ser	Leu	Ala	Ile	Ser	Gly	Leu	Gln
65					70					75					80
Ser	Glu	Asp	Glu	Ala	Asp	Tyr	Asn	Cys	Ala	Ala	Trp	Asp	Asp	Ser	Leu
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Ser	Gly	Pro	Val	Phe	Gly	Gly	Gly	Thr	Lys	Leu	Thr	Val	Leu	Gly	
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1				5					10					15	
Asp	Gln	Val	Leu	Phe	Ile	Asp	Gln	Gly	Asn	Arg	Pro	Leu	Phe	Glu	Asp
			20					25					30		
Met	Thr	Asp	Ser	Asp	Cys	Arg	Asp	Asn	Ala	Pro	Arg	Thr	Ile	Phe	Ile
		35					40					45			
Ile	Ser	Met	Tyr	Lys	Asp	Ser	Gln	Pro	Arg	Gly	Met	Ala	Val	Thr	Ile
	50						55				60				

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20 25 30
Glu Asp